

SEQUENCE LISTING

JUL 6 5 2001

TECH JENTER 1800, 2900

(1) GENERAL INFORMATION:

(i) APPLICANT:

Bahija Jallal Gregory D. Plowman

(ii) TITLE OF INVENTION:

DIAGNOSIS AND TREATMENT OF PTP04 RELATED DISORDERS

(iii) NUMBER OF SEQUENCES:

18

(iv) CORRESPONDENCE ADDRESS:

ADDRESSEE: (A)

Lyon & Lyon

(B) STREET:

633 West Fifth Street

Suite 4700

(C) CITY:

Los Angeles

STATE: (D)

California

COUNTRY: (E)

U.S.A.

ZIP: (F)

90071-2066

(v) COMPUTER READABLE FORM:

MEDIUM TYPE: (A)

3.5" Diskette, 1.44 Mb

storage

(B) COMPUTER:

IBM Compatible

(C) OPERATING SYSTEM: IBM P.C. DOS 5.0

SOFTWARE: (D)

FastSEQ for Windows 2.0

(vi) CURRENT APPLICATION DATA:

(A) APPLICATION NUMBER: To Be Assigned

(B) FILING DATE:

Herewith

(C) CLASSIFICATION:

(vii) PRIOR APPLICATION DATA:

(A) APPLICATION NUMBER: 60/047,222 (B) FILING DATE: May 20, 1997

(B) FILING DATE:

(viii) ATTORNEY/AGENT INFORMATION:

(A) NAME:

Warburg, Richard J.

(B) REGISTRATION NUMBER:

32,327

(C) REFERENCE/DOCKET NUMBER: 234/253

(ix) TELECOMMUNICATION INFORMATION:

(A) TELEPHONE: (213) 489-1600 (B) TELEFAX: (213) 955-0440 (C) TELEX: 67-3510

(2) INFORMATION FOR SEQ ID NO: 1:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 3580 base pairs
(B) TYPE: nucleic acid
(C) STRANDEDNESS: single
(D) TOPOLOGY: linear

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 1:

CCCGGGTGCC CTCCCTCAAC CTACTTATAG ACTATTTTC TTGCTCTGCA GCATGGACCA 60 AAGAGAAATT CTGCAGAAGT TCCTGGATGA GGCCCAAAGC AAGAAAATTA CTAAAGAGGA GTTTGCCAAT GAATTTCTGA AGCTGAAAAG GCAATCTACC AAGTACAAGG CAGACAAAAC 180 CTATCCTACA ACTGTGGCTG AGAAGCCCAA GAATATCAAG AAAAACAGAT ATAAGGATAT TTTGCCCTAT GATTATAGCC GGGTAGAACT ATCCCTGATA ACCTCTGATG AGGATTCCAG CTACATCAAT GCCAACTTCA TTAAGGGAGT TTATGGACCC AAGGCTTATA TTGCCACCCA GGGTCCTTTA TCTACAACCC TCCTGGACTT CTGGAGGATG ATTTGGGAAT ATAGTGTCCT 420 TATCATTGTT ATGGCATGCA TGGAGTATGA AATGGGAAAG AAAAAGTGTG AGCGCTACTG 480 GGCTGAGCCA GGAGAGATGC AGCTGGAATT TGGCCCTTTC TCTGTATCCT GTGAAGCTGA AAAAAGGAAA TCTGATTATA TAATCAGGAC TCTAAAAGTT AAGTTCAATA GTGAAACTCG 600 AACTATCTAC CAGTTTCATT ACAAGAATTG GCCAGACCAT GATGTACCTT CATCTATAGA 660 CCCTATTCTT GAGCTCATCT GGGATGTACG TTGTTACCAA GAGGATGACA GTGTTCCCAT 720 ATGCATTCAC TGCAGTGCTG GCTGTGGAAG GACTGGTGTT ATTTGTGCTA TTGATTATAC ATGGATGTTG CTAAAAGATG GGATAATTCC TGAGAACTTC AGTGTTTTCA GTTTGATCCG 840 GGAAATGCGG ACACAGAGGC CTTCATTAGT TCAAACGCAG GAACAATATG AACTGGTCTA 900 CAATGCTGTA TTAGAACTAT TTAAGAGACA GATGGATGTT ATCAGAGATA AACATTCTGG 960 AACAGAGAGT CAAGCAAAGC ATTGTATTCC TGAGAAAAAT CACACTCTCC AAGCAGACTC TTATTCTCCT AATTTACCAA AAAGTACCAC AAAAGCAGCA AAAATGATGA ACCAACAAAG 1080 GACAAAAATG GAAATCAAAG AATCTTCTTC CTTTGACTTT AGGACTTCTG AAATAAGTGC 1140 AAAAGAAGAG CTAGTTTTGC ACCCTGCTAA ATCAAGCACT TCTTTTGACT TTCTGGAGCT 1200 AAATTACAGT TTTGACAAAA ATGCTGACAC AACCATGAAA TGGCAGACAA AGGCATTTCC 1260 AATAGTTGGG GAGCCTCTTC AGAAGCATCA AAGTTTGGAT TTGGGCTCTC TTTTGTTTGA 1320 GGGATGTTCT AATTCTAAAC CTGTAAATGC AGCAGGAAGA TATTTTAATT CAAAGGTGCC 1380 AATAACACGG ACCAAATCAA CTCCTTTTGA ATTGATACAG CAGAGAGAAA CCAAGGAGGT 1440 GGACAGCAAG GAAAACTTTT CTTATTTGGA ATCTCAACCA CATGATTCTT GTTTTGTAGA 1500 GATGCAGGCT CAAAAAGTAA TGCATGTTTC TTCAGCAGAA CTGAATTATT CACTGCCATA 1560 TGACTCTAAA CACCAAATAC GTAATGCCTC TAATGTAAAG CACCATGACT CTAGTGCTCT 1620 TGGTGTATAT TCTTACATAC CTTTAGTGGA AAATCCTTAT TTTTCATCAT GGCCTCCAAG 1680 TGGTACCAGT TCTAAGATGT CTCTTGATTT ACCTGAGAAG CAAGATGGAA CTGTTTTCC 1740 TTCTTCTCTG TTGCCAACAT CCTCTACATC CCTCTTCTCT TATTACAATT CACATGATTC 1800 TTTATCACTG AATTCTCCAA CCAATATTTC CTCACTATTG AACCAGGAGT CAGCTGTACT 1860 AGCAACTGCT CCAAGGATAG ATGATGAAAT CCCCCCTCCA CTTCCTGTAC GGACACCTGA 1920 ATCATTTATT GTGGTTGAGG AAGCTGGAGA ATTCTCACCA AATGTTCCCA AATCCTTATC 1980 CTCAGCTGTG AAGGTAAAAA TTGGAACATC ACTGGAATGG GGTGGAACAT CTGAACCAAA 2040 GAAATTTGAT GACTCTGTGA TACTTAGACC AAGCAAGAGT GTAAAACTCC GAAGTCCTAA 2100 ATCAGAACTA CATCAAGATC GTTCTTCTCC CCCACCTCCT CTCCCAGAAA GAACTCTAGA GTCCTTCTTT CTTGCCGATG AAGATTGTAT GCAGGCCCAA TCTATAGAAA CATATTCTAC TAGCTATCCT GACACCATGG AAAATTCAAC ATCTTCAAAA CAGACACTGA AGACTCCTGG 2280 AAAAAGTTTC ACAAGGAGTA AGAGTTTGAA AATTTTGCGA AACATGAAAA AGAGTATCTG 2340 TAATTCTTGC CCACCAAACA AGCCTGCAGA ATCTGTTCAG TCAAATAACT CCAGCTCATT 2400 TCTGAATTTT GGTTTTGCAA ACCGTTTTTC AAAACCCAAA GGACCAAGGA ATCCACCACC 2460 AACTTGGAAT ATTTAATAAA ACTCCAGATT TATAATAATA TGGGCTGCAA GTACACCTGC 2520 AAATAAAACT ACTAGAATAC TGCTAGTTAA AATAAGTGCT CTATATGCAT AATATCAAAT ATGAAGATAT GCTAATGTGT TAATAGCTTT TAAAAGAAAA GCAAAATGCC AATAAGTGCC 2640 AGTTTTGCAT TTTCATATCA TTTGCATTGA GTTGAAAACT GCAAATAAAA GTTTGTCACT 2700 TGAGCTTATG TACAGAATGC TATATGAGAA ACACTTTTAG AATGGATTTA TTTTTCATTT 2760 TTGCCAGTTA TTTTATTTT CTTTTACTTT TTTACATAAA CATAAACTTC AAAAGGTTTG 2820 TAAGATTTGG ATCTCAACTA ATTTCTACAT TGCCAGAATA TACTATAAAA AGTTAAAAAA 2880 AAACTTACTT TGTGGGTTGC AATACAAACT GCTCTTGACA ATGACTATTC CCTGACAGTT 2940 ATTTTTGCCT AAATGGAGTA TACCTTGTAA ATCTTCCCAA ATGTTGTGGA AAACTGGAAT 3000 ATTAAGAAAA TGAGAAATTA TATTTATTAG AATAAAATGT GCAAATAATG ACAATTATTT 3060 GAATGTAACA AGGAATTCAA CTGAAATCCT GATAAGTTTT AACCAAAGTC ATTAAATTAC 3120 CAATTCTAGA AAAGTAATCA ATGAAATATA ATAGCTATCT TTTGGTAGCA AAAGATATAA 3180 ATTGTATATG TTTATACAGG ATCTTTCAGA TCATGTGCAA TTTTTATCTA ACCAATCAGA 3240 AATACTAGTT TAAAATGAAT TTCTATATGA ATATGGATCT GCCATAAGAA AATCTAGTTC 3300 AACTCTAATT TTATGTAGTA AATAAATTGG CAGGTAATTG TTTTTACAAA GAATCCACCT 3360 GACTTCCCCT AATGCATTAA AAATATTTTT ATTTAAATAA CTTTATTTAT AACTTTTAGA AACATGTAGT ATTGTTTAAA CATCATTTGT TCTTCAGTAT TTTTCATTTG GAAGTCCAAT 3480 AGGGCAAATT GAATGAAGTA TTATTATCTG TCTCTTGTAG TACAATGTAT CCAACAGACA 3540 3580

(2) INFORMATION FOR SEQ ID NO: 2:

(i) SEQUENCE CHARACTERISTICS:

(A) LENGTH: 807 amino acids
(B) TYPE: amino acid
(C) STRANDEDNESS: single

(D) TOPOLOGY: linear

(ii) MOLECULE TYPE: peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 2:

Met Asp Gln Arg Glu Ile Leu Gln Lys Phe Leu Asp Glu Ala Gln Ser 1 10 15

Lys Lys Ile Thr Lys Glu Glu Phe Ala Asn Glu Phe Leu Lys Leu Lys 20 25 30

Arg Gln Ser Thr Lys Tyr Lys Ala Asp Lys Thr Tyr Pro Thr Thr Val 40 Ala Glu Lys Pro Lys Asn Ile Lys Lys Asn Arg Tyr Lys Asp Ile Leu Pro Tyr Asp Tyr Ser Arg Val Glu Leu Ser Leu Ile Thr Ser Asp Glu 75 Asp Ser Ser Tyr Ile Asn Ala Asn Phe Ile Lys Gly Val Tyr Gly Pro Lys Ala Tyr Ile Ala Thr Gln Gly Pro Leu Ser Thr Thr Leu Leu Asp 100 Phe Trp Arg Met Ile Trp Glu Tyr Ser Val Leu Ile Ile Val Met Ala 120 Cys Met Glu Tyr Glu Met Gly Lys Lys Lys Cys Glu Arg Tyr Trp Ala 135 Glu Pro Gly Glu Met Gln Leu Glu Phe Gly Pro Phe Ser Val Ser Cys 150 145 Glu Ala Glu Lys Arg Lys Ser Asp Tyr Ile Ile Arg Thr Leu Lys Val 170 Lys Phe Asn Ser Glu Thr Arg Thr Ile Tyr Gln Phe His Tyr Lys Asn 180 Trp Pro Asp His Asp Val Pro Ser Ser Ile Asp Pro Ile Leu Glu Leu Ile Trp Asp Val Arg Cys Tyr Gln Glu Asp Asp Ser Val Pro Ile Cys Ile His Cys Ser Ala Gly Cys Gly Arg Thr Gly Val Ile Cys Ala Ile 225 Asp Tyr Thr Trp Met Leu Leu Lys Asp Gly Ile Ile Pro Glu Asn Phe 250 Ser Val Phe Ser Leu Ile Arg Glu Met Arg Thr Gln Arg Pro Ser Leu 260 Val Gln Thr Gln Glu Gln Tyr Glu Leu Val Tyr Asn Ala Val Leu Glu 275

Leu Phe Lys Arg Gln Met Asp Val Ile Arg Asp Lys His Ser Gly Thr

295

300

290

Glu 8	Ser	Gln ·	Ala	Lys	His 310	Cys	Ile :	Pro	Glu	Lys 315	Asn	His	Thr I	Leu C	Hln 320
Ala .	Asp	Ser	Tyr	Ser 325	Pro	Asn	Leu	Pro	Lys 330	Ser	Thr	Thr	Lys :	Ala <i>l</i> 335	Ala
Lys	Met	Met	Asn 340	Gln	Gln	Arg	Thr	Lys 345	Met	Glu	Ile	Lys	Glu 350	Ser S	Ser
		355					360				Lys	505			
	370					375					Phe 380				
385					390					333					
				405					410		His				
			420					425			. Ser				
		435	5				440				Ile				
	450	1				455					1 Thr 460	,			
465	i				470)				- <u></u> / .					
				485	5				49	U	s Val				
			50	0				50:	>				-		Ala
		51	.5				540	J			u Gl	•	_		
Il	e Pr 53		u Va	l Gl	u As:	n Pro	о Ту: 5	r Ph	e Se	r Se	r Tr	p Pr O	o Pro	o Ser	: Gly
Th 54		r Se	er Ly	rs Me	t Se 55	r Le	u As	p Le	u Pr	o Gl 59	u Ly 55	s Gl	n As	p Gly	7 Thr 560
۷a	l Ph	ie Pi	co Se	er Se	r Le	u Le	u Pr	o Th	ır Se	er Se 70	er Th	ır Se	r Le	u Ph	e Ser 5

Tyr Tyr Asn Ser His Asp Ser Leu Ser Leu Asn Ser Pro Thr Asn Ile
580 585 590

Ser Ser Leu Leu Asn Gln Glu Ser Ala Val Leu Ala Thr Ala Pro Arg 595 600 605

Ile Asp Asp Glu Ile Pro Pro Pro Leu Pro Val Arg Thr Pro Glu Ser 610 620

Phe Ile Val Val Glu Glu Ala Gly Glu Phe Ser Pro Asn Val Pro Lys 625 630 635

Ser Leu Ser Ser Ala Val Lys Val Lys Ile Gly Thr Ser Leu Glu Trp 645 650 655

Gly Gly Thr Ser Glu Pro Lys Lys Phe Asp Asp Ser Val Ile Leu Arg 660 665 670

Pro Ser Lys Ser Val Lys Leu Arg Ser Pro Lys Ser Glu Leu His Gln 675 680

Asp Arg Ser Ser Pro Pro Pro Pro Leu Pro Glu Arg Thr Leu Glu Ser 690 695 700

Phe Phe Leu Ala Asp Glu Asp Cys Met Gln Ala Gln Ser Ile Glu Thr 705 710 715 720

Tyr Ser Thr Ser Tyr Pro Asp Thr Met Glu Asn Ser Thr Ser Ser Lys 725 730 735

Gln Thr Leu Lys Thr Pro Gly Lys Ser Phe Thr Arg Ser Lys Ser Leu 740 745 750

Lys Ile Leu Arg Asn Met Lys Lys Ser Ile Cys Asn Ser Cys Pro Pro 755 760 765

Asn Lys Pro Ala Glu Ser Val Gln Ser Asn Asn Ser Ser Ser Phe Leu 770 780

Asn Phe Gly Phe Ala Asn Arg Phe Ser Lys Pro Lys Gly Pro Arg Asn 785 790 795 800

Pro Pro Pro Thr Trp Asn Ile 805 (2) INFORMATION FOR SEQ ID NO: 3: (i) SEQUENCE CHARACTERISTICS: 23 base pairs (A) LENGTH: nucleic acid (B) TYPE: single (C) STRANDEDNESS: linear (D) TOPOLOGY: (ix) FEATURE: (D) OTHER INFORMATION: The letter "Y" stands for C or T. The letter "V" stands for A, C or The letter "R" stands for A or G. The letter "N" stands for A, C, G or T. (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 3: 23 GAYTTYTGGV RNATGRTNTG GGA (2) INFORMATION FOR SEQ ID NO: 4: (i) SEQUENCE CHARACTERISTICS: 23 base pairs (A) LENGTH: nucleic acid (B) TYPE: single (C) STRANDEDNESS: (D) TOPOLOGY: linear (ix) FEATURE: The letter "S" stands for C or G. (D) OTHER INFORMATION: The letter "Y" stands for C or T. The letter "N" stands for A, C, G or T. The letter "W" stands for A or T. The letter "R" stands for A or G.

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(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 4:

CGGCCSAYNC CNGCNSWRCA RTG

- (2) INFORMATION FOR SEQ ID NO: 5:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH:

8 amino acids amino acid

(B) TYPE:

(C) STRANDEDNESS: single

(D) TOPOLOGY:

(ii) MOLECULE TYPE: peptide

- (ix) FEATURE:
 - (D) OTHER INFORMATION:

"Xaa" in positions 4 and 6 stand for an unspecified amino acid.

"Xaa" in position 8 stands for

either Glu or Asp.

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 5:

Asp Phe Trp Xaa Met Xaa Trp Xaa

- (2) INFORMATION FOR SEQ ID NO: 6:
 - (i) SEQUENCE CHARACTERISTICS:

(A) LENGTH:

7 amino acids

(B) TYPE:

amino acid

(C) STRANDEDNESS: single

(D) TOPOLOGY:

linear

(ii) MOLECULE TYPE: peptide

- (ix) FEATURE:

(D) OTHER INFORMATION: "Xaa" in positions 3 and 6 stand

for an unspecified amino acid.

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 6:

His Cys Xaa Ala Gly Xaa Gly

5

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(2) INFO	RMATION FOR SEQ ID NO:	7:	
(i)	SEQUENCE CHARACTERISTIC	S:	
	(A) LENGTH: (B) TYPE: (C) STRANDEDNESS: (D) TOPOLOGY:	34 base pairs nucleic acid single linear	
(xi)	SEQUENCE DESCRIPTION: S	SEQ ID NO: 7:	
CACCGTT	CGA GTATTTCAGA TTGTGAAGA	A GTCC	34
(2) INF	ORMATION FOR SEQ ID NO:	8:	
(i)	SEQUENCE CHARACTERISTI	CS:	
	(A) LENGTH:(B) TYPE:(C) STRANDEDNESS:(D) TOPOLOGY:	34 base pairs nucleic acid single linear	
(xi)	SEQUENCE DESCRIPTION:	SEQ ID NO: 8:	
GGACTTC	TTC ACAATCTGAA ATACTCGAA	C GGTG	34
(2) INE	FORMATION FOR SEQ ID NO:	9:	
(i)	SEQUENCE CHARACTERIST	CCS:	
	(A) LENGTH: (B) TYPE: (C) STRANDEDNESS: (D) TOPOLOGY:	33 base pairs nucleic acid single linear	
(xi) SEQUENCE DESCRIPTION:	SEQ ID NO: 9:	
CCGTTA	TGTG AGGAAGAGCC ACATTACA	GG ACC	33
(2) IN	FORMATION FOR SEQ ID NO:	10:	
(i) SEQUENCE CHARACTERIST	PICS:	
	(A) LENGTH:(B) TYPE:(C) STRANDEDNESS:(D) TOPOLOGY:	33 base pairs nucleic acid single linear	

	(xi)	SEQUENCE DESCRIPTION: SEQ	ID NO: 10:					
GGT	CCTGT	LA TGTGGCTCTT CCTCACATAA CC	GG	33				
(2)	INFO	RMATION FOR SEQ ID NO: 11	:					
	(i)	SEQUENCE CHARACTERISTICS:						
		(A) LENGTH: (B) TYPE: (C) STRANDEDNESS: (D) TOPOLOGY:	23 base pairs nucleic acid single linear					
	(xi)	SEQUENCE DESCRIPTION: SEQ	ID NO: 11:					
GG	CATGCA	TG GAGTATGAAA TGG	•	23				
(2)	INFC	RMATION FOR SEQ ID NO: 12	•					
	(i)	SEQUENCE CHARACTERISTICS:						
		(A) LENGTH: (B) TYPE: (C) STRANDEDNESS: (D) TOPOLOGY:	30 base pairs nucleic acid single linear					
	(xi)	SEQUENCE DESCRIPTION: SEQ) ID NO: 12:					
CG	TACATO	CC AGATGAGCTC AAGAATAGGG		30				
(2) INFORMATION FOR SEQ ID NO: 13:								
	(i)	SEQUENCE CHARACTERISTICS	:					
		(A) LENGTH:(B) TYPE:(C) STRANDEDNESS:(D) TOPOLOGY:	10 amino acids amino acid single linear					
	(ii)	MOLECULE TYPE:	peptide					
	(xi)	SEQUENCE DESCRIPTION: SE	Q ID NO: 13:					
Т	yr Pro	Tyr Asp Val Pro Asp Tyr A	la Ser 10					

(i) SEQUENCE CHARACTERISTICS: 5 amino acids (A) LENGTH: amino acid single (B) TYPE: (C) STRANDEDNESS: linear (D) TOPOLOGY: peptide (ii) MOLECULE TYPE: (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 14: His Cys Ser Ala Gly (2) INFORMATION FOR SEQ ID NO: 15: (i) SEQUENCE CHARACTERISTICS: 31 amino acids (A) LENGTH: amino acid (B) TYPE: single (C) STRANDEDNESS: linear (D) TOPOLOGY: (ii) MOLECULE TYPE: peptide (xi) SEQUENCE DESCRIPTION: SEQ ID NO: 15: Ser Trp Pro Pro Ser Gly Thr Ser Ser Lys Met Ser Leu Asp Asp Leu Pro Glu Lys Gln Asp Gly Thr Val Phe Pro Ser Ser Leu Leu Pro 20 (2) INFORMATION FOR SEQ ID NO: 16: (i) SEQUENCE CHARACTERISTICS: 30 amino acids (A) LENGTH: amino acid (B) TYPE: single (C) STRANDEDNESS: linear (D) TOPOLOGY: peptide (ii) MOLECULE TYPE:

(2) INFORMATION FOR SEQ ID NO: 14:

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: . 16:

Tyr Ser Leu Pro Tyr Asp Ser Lys His Gln Ile Arg Asn Ala Ser Asn

Val Lys His His Asp Ser Ser Ala Leu Gly Val Tyr Ser Tyr

- (2) INFORMATION FOR SEQ ID NO: 17:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH:

30 amino acids

- (B) TYPE:
- amino acid
- (C) STRANDEDNESS: single
- (D) TOPOLOGY:
- (ii) MOLECULE TYPE:

peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 17:

His Thr Leu Gln Ala Asp Ser Tyr Ser Pro Asn Leu Pro Lys Ser Thr

Thr Lys Ala Ala Lys Met Met Asn Gln Gln Arg Thr Lys Cys 20

- (2) INFORMATION FOR SEQ ID NO: 18:
 - (i) SEQUENCE CHARACTERISTICS:
 - (A) LENGTH:

802 amino acids

(B) TYPE:

amino acid

(C) STRANDEDNESS:

single

(D) TOPOLOGY:

linear

(ii) MOLECULE TYPE:

peptide

(xi) SEQUENCE DESCRIPTION: SEQ ID NO: 18:

Met Asp Gln Arg Glu Ile Leu Gln Gln Leu Leu Lys Glu Ala Gln Lys

Lys Lys Leu Asn Ser Glu Glu Phe Ala Ser Glu Phe Leu Lys Leu Lys

Arg Gln Ser Thr Lys Tyr Lys Ala Asp Lys Ile Tyr Pro Thr Thr Val 40 Ala Gln Arg Pro Lys Asn Ile Lys Lys Asn Arg Tyr Lys Asp Ile Leu Pro Tyr Asp His Ser Leu Val Glu Leu Ser Leu Leu Thr Ser Asp Glu Asp Ser Ser Tyr Ile Asn Ala Ser Phe Ile Lys Gly Val Tyr Gly Pro 90 85 Lys Ala Tyr Ile Ala Thr Gln Gly Pro Leu Ser Thr Thr Leu Leu Asp 105 Phe Trp Arg Met Ile Trp Glu Tyr Arg Ile Leu Val Ile Val Met Ala 120 Cys Met Glu Phe Glu Met Gly Lys Lys Lys Cys Glu Arg Tyr Trp Ala Glu Pro Gly Glu Thr Gln Leu Gln Phe Gly Pro Phe Ser Ile Ser Cys 155 150 Glu Ala Glu Lys Lys Lys Ser Asp Tyr Lys Ile Arg Thr Leu Lys Ala 170 165 Lys Phe Asn Asn Glu Thr Arg Ile Ile Tyr Gln Phe His Tyr Lys Asn 185 180 Trp Pro Asp His Asp Val Pro Ser Ser Ile Asp Pro Ile Leu Gln Leu 200 Ile Trp Asp Met Arg Cys Tyr Gln Glu Asp Asp Cys Val Pro Ile Cys Ile His Cys Ser Ala Gly Cys Gly Arg Thr Gly Val Ile Cys Ala Val 230 Asp Tyr Thr Trp Met Leu Leu Lys Asp Gly Ile Ile Pro Lys Asn Phe 250 Ser Val Phe Asn Leu Ile Gln Glu Met Arg Thr Gln Arg Pro Ser Leu 265 260 Val Gln Thr Gln Glu Gln Tyr Glu Leu Val Tyr Ser Ala Val Leu Glu 280 Leu Phe Lys Arg His Met Asp Val Ile Ser Asp Asn His Leu Gly Arg 295

Glu Ile Gln Ala Gln Cys Ser Ile Pro Glu Gln Ser Leu Thr Val Glu 310 305 Ala Asp Ser Cys Pro Leu Asp Leu Pro Lys Asn Ala Met Arg Asp Val 330 325 Lys Thr Thr Asn Gln His Ser Lys Gln Gly Ala Glu Ala Glu Ser Thr 345 Gly Gly Ser Ser Leu Gly Leu Arg Thr Ser Thr Met Asn Ala Glu Glu Glu Leu Val Leu His Ser Ala Lys Ser Ser Pro Ser Phe Asn Cys Leu 375 Glu Leu Asn Cys Gly Cys Asn Asn Lys Ala Val Ile Thr Arg Asn Gly Gln Ala Arg Ala Ser Pro Val Val Gly Glu Pro Leu Gln Lys Tyr Gln Ser Leu Asp Phe Gly Ser Met Leu Phe Gly Ser Cys Pro Ser Ala Leu 425 Pro Ile Asn Thr Ala Asp Arg Tyr His Asn Ser Lys Gly Pro Val Lys Arg Thr Lys Ser Thr Pro Phe Glu Leu Ile Gln Gln Arg Lys Thr Asn 455 Asp Leu Ala Val Gly Asp Gly Phe Ser Cys Leu Glu Ser Gln Leu His Glu His Tyr Ser Leu Arg Glu Leu Gln Val Gln Arg Val Ala His Val Ser Ser Glu Glu Leu Asn Tyr Ser Leu Pro Gly Ala Cys Asp Ala Ser 500 Cys Val Pro Arg His Ser Pro Gly Ala Leu Arg Val His Leu Tyr Thr Ser Leu Ala Glu Asp Pro Tyr Phe Ser Ser Pro Pro Asn Ser Ala Asp Ser Lys Met Ser Phe Asp Leu Pro Glu Lys Gln Asp Gly Ala Thr 555 550 545 Ser Pro Gly Ala Leu Leu Pro Ala Ser Ser Thr Thr Ser Phe Phe Tyr

570

565

Ser Asn Pro His Asp Ser Leu Val Met Asn Thr Leu Thr Ser Phe Ser 585 580 Pro Pro Leu Asn Gln Glu Thr Ala Val Glu Ala Pro Ser Arg Arg Thr 600 Asp Asp Glu Ile Pro Pro Pro Leu Pro Glu Arg Thr Pro Glu Ser Phe 610 Ile Val Val Glu Glu Ala Gly Glu Pro Ser Pro Arg Val Thr Glu Ser 635 Leu Pro Leu Val Val Thr Phe Gly Ala Ser Pro Glu Cys Ser Gly Thr 650 645 Ser Glu Met Lys Ser His Asp Ser Val Gly Phe Thr Pro Ser Lys Asn 665 Val Lys Leu Arg Ser Pro Lys Ser Asp Arg His Gln Asp Gly Ser Pro 680 Pro Pro Pro Leu Pro Glu Arg Thr Leu Glu Ser Phe Phe Leu Ala Asp Glu Asp Cys Ile Gln Ala Gln Ala Val Gln Thr Ser Ser Thr Ser Tyr 715 710 Pro Glu Thr Thr Glu Asn Ser Thr Ser Ser Lys Gln Thr Leu Arg Thr 725 Pro Gly Lys Ser Phe Thr Arg Ser Lys Ser Leu Lys Ile Phe Arg Asn 745 740 Met Lys Lys Ser Val Cys Asn Ser Ser Ser Pro Ser Lys Pro Thr Glu 760 Arg Val Gln Pro Lys Asn Ser Ser Ser Phe Leu Asn Phe Gly Phe Gly 775 770 Asn Arg Phe Ser Lys Pro Lys Gly Pro Arg Asn Pro Pro Ser Ala Trp 795 790

Asn Met